HIGHLY SPECIFIC MODULATORS OF GTPASES FOR TARGET VALIDATION Kavita Shah, et al. Serial No.: Attorney Docket No.: 36-001720US

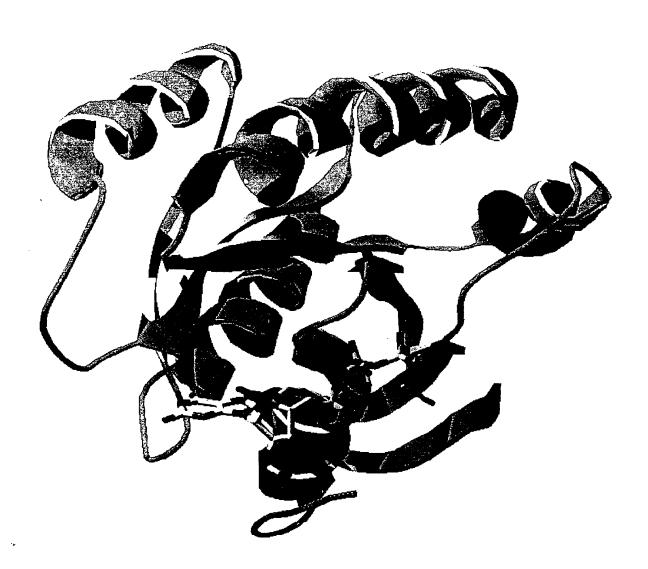


Fig. 1

HIGHLY SPECIFIC MODULATORS OF GTPASES FOR TARGET VALIDATION Kavita Shah, et al.

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$$R_1$$
 N
 N
 N
 N
 N
 N
 N
 N

Fig. 2A

Fig. 2B

Fig. 2C

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Fig. 2D
$$\underset{\text{Ho}-P-O-P-O}{\overset{\text{R}}{\underset{\text{NH}_2}{\longrightarrow}}}$$

Fig. 2F

Fig. 2G

Fig. 2H

Fig. 2I



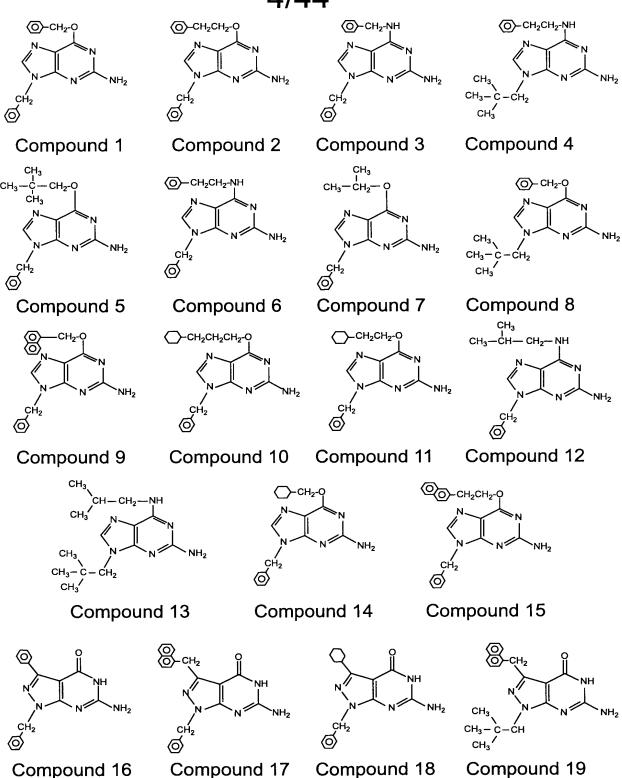


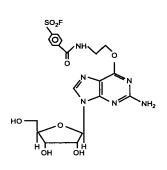
Fig. 3

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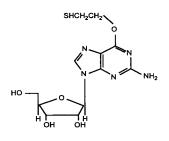
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Compound 21



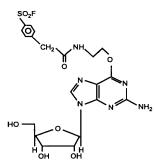
Compound 24



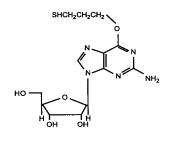
Compound 27

Compound 30

Compound 22



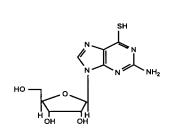
Compound 25



Compound 28

Compound 31

Compound 23



Compound 26

Compound 29

Compound 32

Fig. 4

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Fig. 4 Cont.

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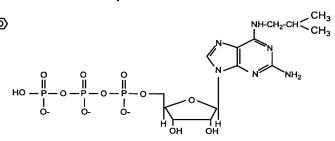
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Compound 39

NH-CH2CH2-O

Compound 40

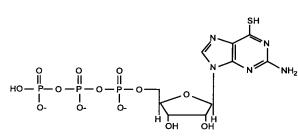


Compound 41

O-CH2-CH=CH2

Compound 42

Compound 43



Compound 44

$$\begin{array}{c} \text{CH}_{3} \\ \text{CH}_{2} \\$$

Compound 46

Compound 47

Compound 48

Fig. 5

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Compound 51

Compound 53

Compound 55

Compound 50

Compound 52

Compound 54

Compound 56

Fig. 5 Cont.

Compound 98

Compound 99

Compound 100

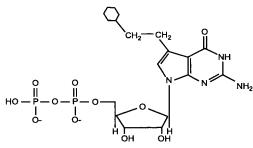
Compound 101

Compound 102

Compound 103

Fig. 5 Cont.

Compound 57



Compound 59

$$\begin{array}{c|c} & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & &$$

Compound 61

Compound 63

Compound 58

$$\begin{array}{c|c} & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & &$$

Compound 60

$$\begin{array}{c|c} \bigcirc CH_2 \\ \hline \\ CH_2 - CH_2 \\ \hline \\ CH_2 - CH_2 \\ \hline \\ N \\ NH_2 \\ \hline \\ NH_2$$

$$\begin{array}{c|c} & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & &$$

Compound 64

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Fig. 6 Cont.

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Compound 65

Compound 67

Compound 69

Compound 71

Compound 73

Compound 66

Compound 68

Compound 72

Compound 74

Fig. 7

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Compound 75

Compound 77

$$CH_3CSCH_2CH_2O - P - O \longrightarrow H OH OH$$

$$CH_3CSCH_2CH_2O + OH OH OH$$

Compound 79

Compound 81

Compound 76

Compound 80

Compound 82

Fig. 7 Cont.

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Compound 88

Compound 89

Compound 90

Fig. 7 Cont.

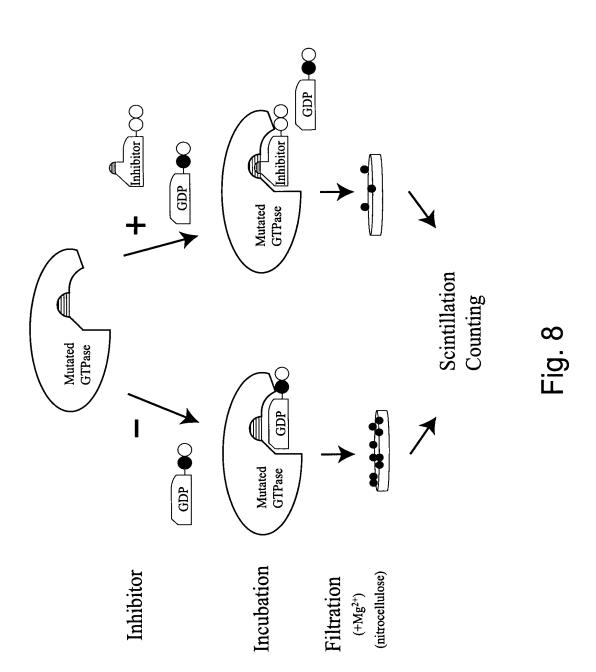


Fig. 9A

Fig. 9B

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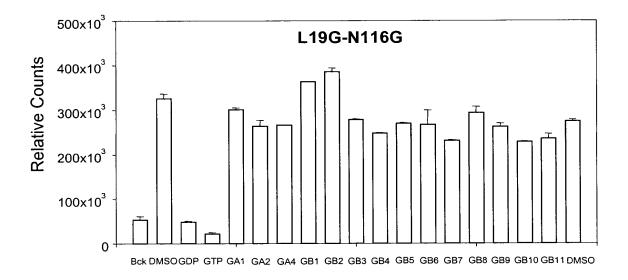


Fig. 10

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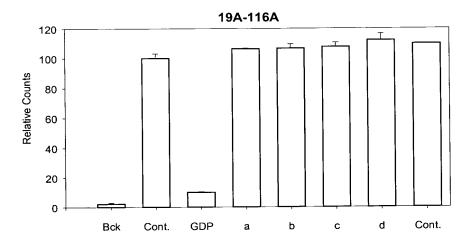


Fig. 11

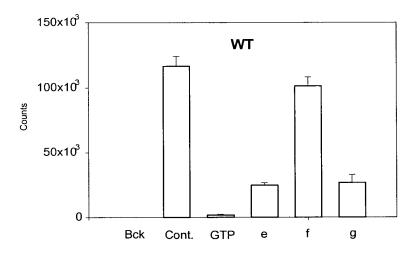


Fig. 12A

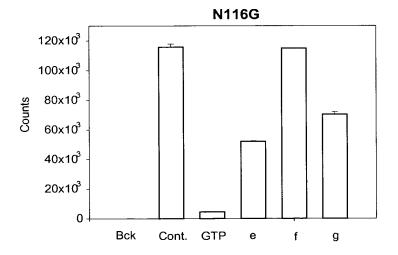
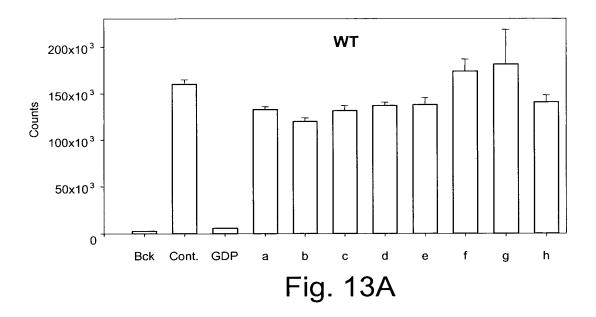


Fig. 12B



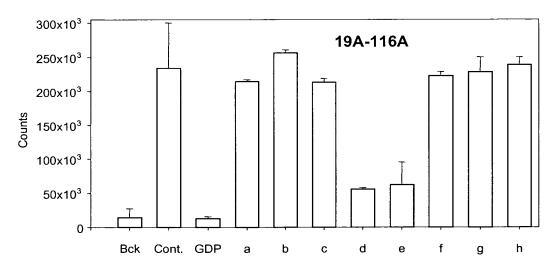


Fig. 13B

HIGHLY SPECIFIC MODULATORS OF GTPASES FOR TARGET VALIDATION Kavita Shah, et al. Serial No.: Attorney Docket No.: 36-001720US

HIGHLY SPECIFIC MODULATORS OF GTPASES FOR TARGET VALIDATION Kavita Shah, et al. Serial No.: Attorney Docket No.: 36-001720US

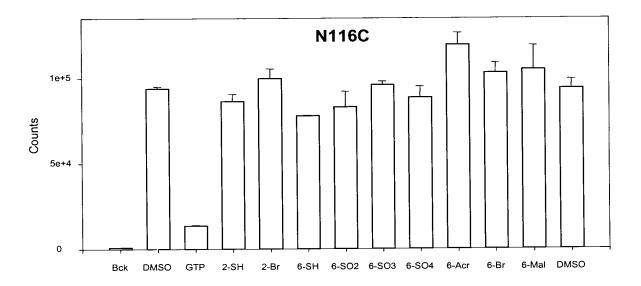


Fig. 15

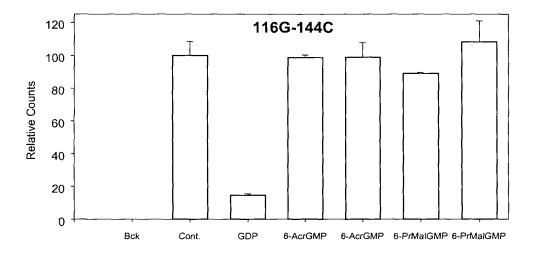


Fig. 16

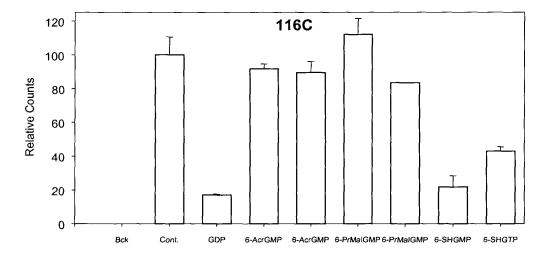
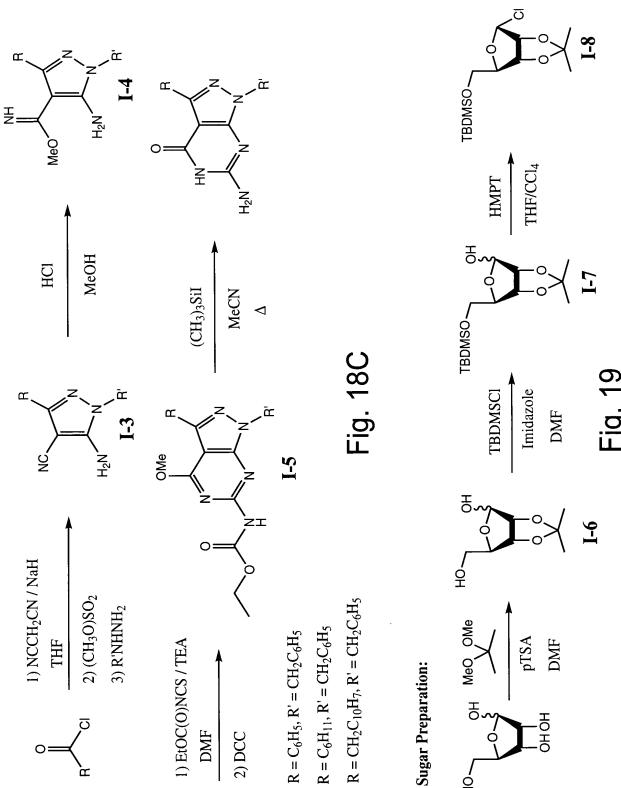


Fig. 17

HIGHLY SPECIFIC MODULATORS OF GTPASES FOR TARGET VALIDATION

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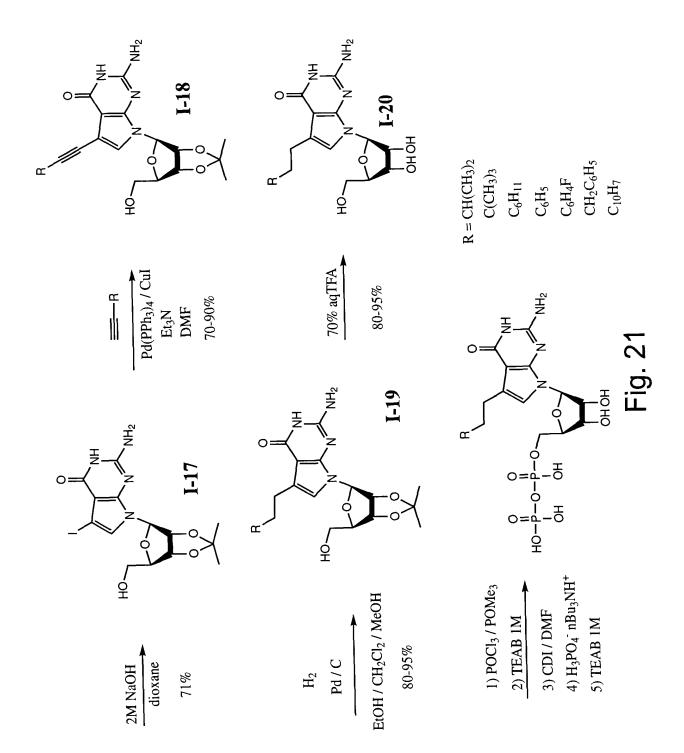
HIGHLY SPECIFIC MODULATORS OF GTPASES FOR TARGET VALIDATION Kavita Shah, et al. Serial No.: Attorney Docket No.: 36-001720US

26/44 TBDMSO~ KOH / TDA-1 MeCN POCl₃ TBDMSO-I-12 1) NaOMe / MeOH Bomb NIS THF 84% 2) Me_2S I-14 Pyridine mCPBA

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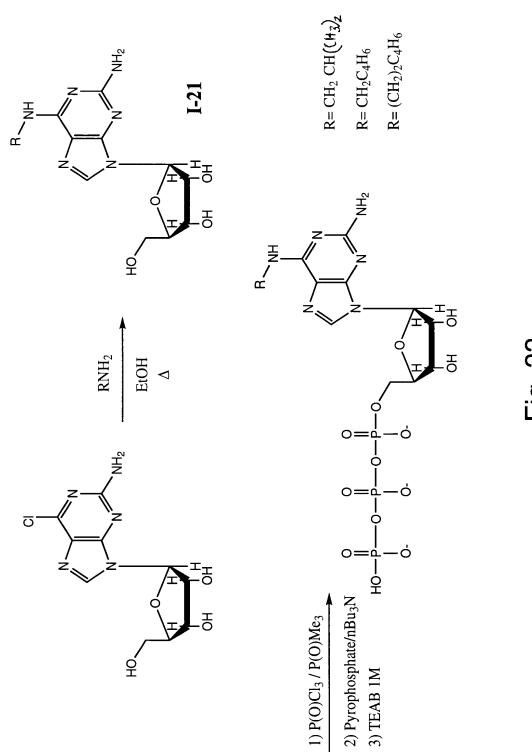
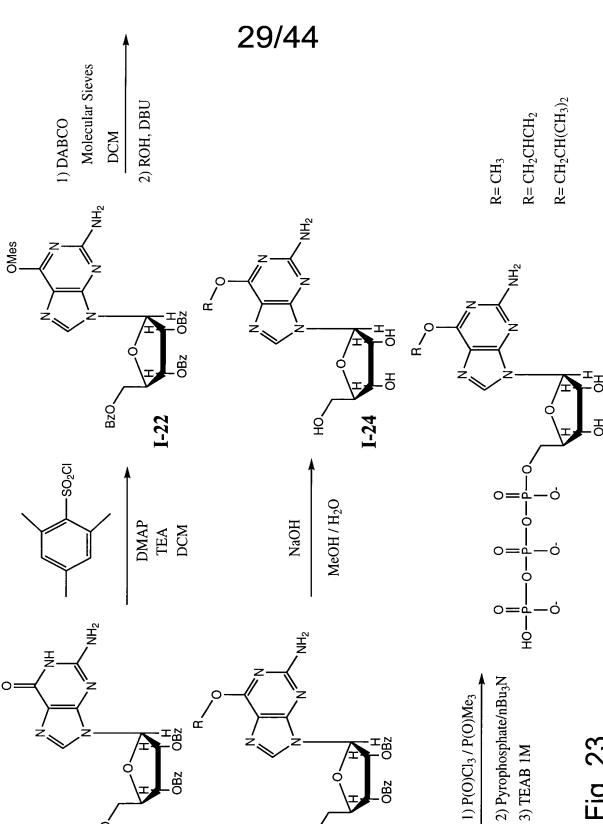
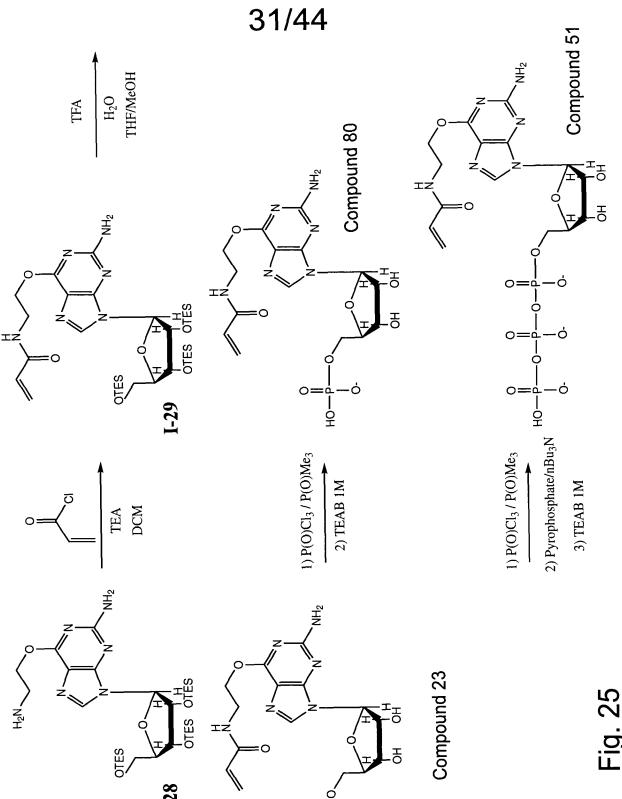


Fig. 22



I-23

BZO



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$$R = (CH2)2SH$$

$$(CH2)3SH$$

$$(CH2)4SH$$

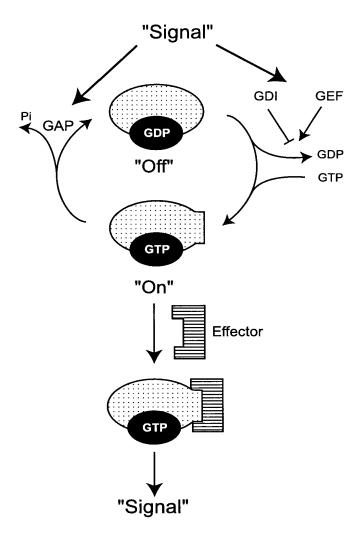


Fig. 32

HIGHLY SPECIFIC MODULATORS OF GTPASES FOR TARGET VALIDATION Kavita Shah, et al. Serial No.: Attorney Docket No.: 36-001720US

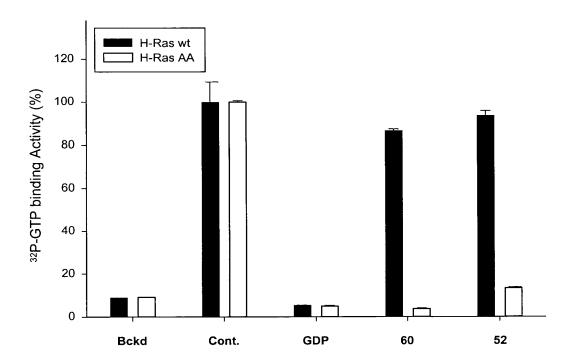


Fig. 33A

HIGHLY SPECIFIC MODULATORS OF GTPASES FOR TARGET VALIDATION
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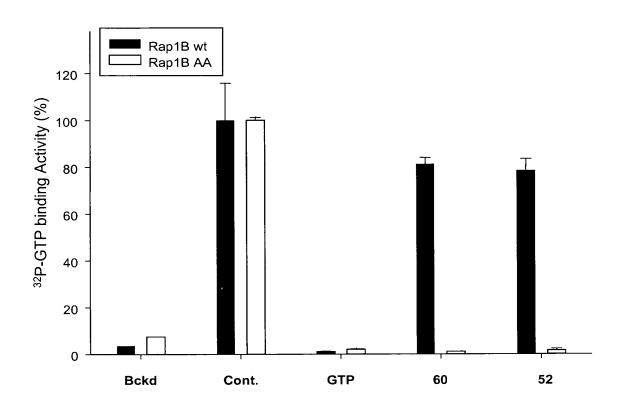


Fig. 33B

HIGHLY SPECIFIC MODULATORS OF GTPASES FOR TARGET VALIDATION Kavita Shah, et al. Serial No.: Attorney Docket No.: 36-001720US

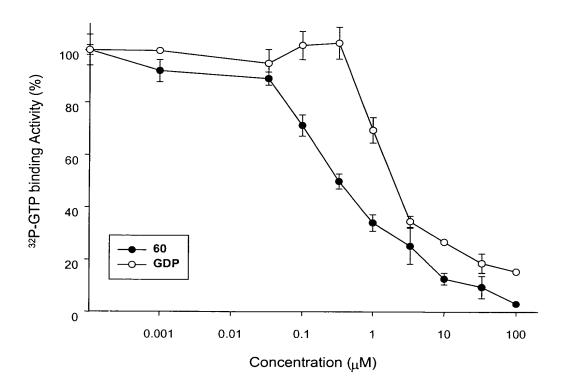


Fig. 34

HIGHLY SPECIFIC MODULATORS OF GTPASES FOR TARGET VALIDATION

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AA	ı	60 52	
AA	ı	09	
AA	GTP	ı	
AA AA AA AA	GDP GTP	ı	
AA	ı	1	
wt	GTP	1	
wt	GDP GTP	1	
Ras	GXP	Compound	H-Ras -

RBD pull-down experiment:

Fig. 35

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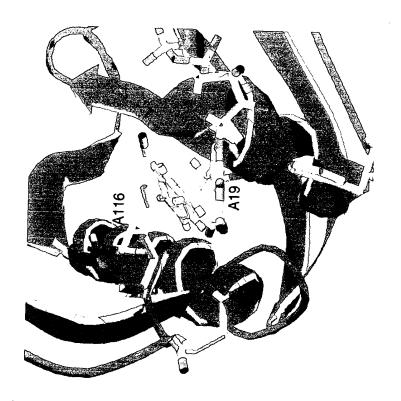


Fig. 36B

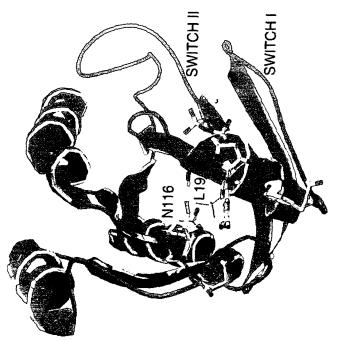


Fig. 36A

HIGHLY SPECIFIC MODULATORS OF GTPASES FOR TARGET VALIDATION Kavita Shah, et al. Serial No.: Attorney Docket No.: 36-001720US

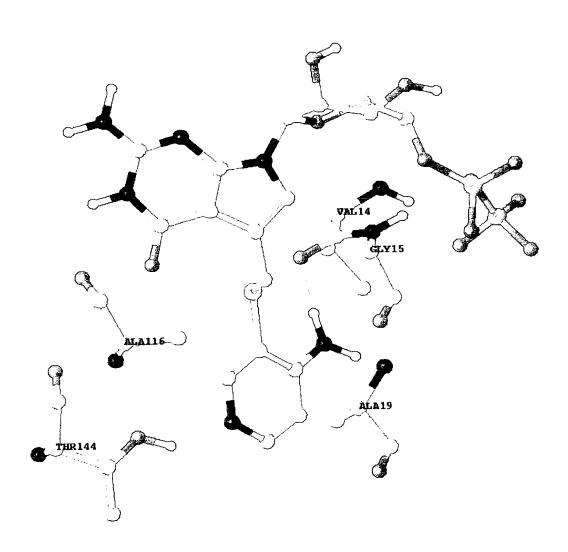


Fig. 37